

**AMENDMENTS****In the Claims**

- 1.(canceled)
- 2.(canceled)
- 3.(canceled)
- 4.(canceled)
- 5.(canceled)
- 6.(canceled)
- 7.(canceled)
- 8.(canceled)
- 9.(canceled)

1      10.(previously presented)    A composition comprising a polymerizing agent including a molecular  
2      and/or atomic tag covalently bonded to a site on the polymerizing agent and a monomer including  
3      a molecular and/or atomic tag, where at least one of the tags has a fluorescence property that  
4      undergoes a change before, during and/or after each of a sequence of monomer incorporations due  
5      to an interaction between the polymerizing agent tag and the monomer tag and where the changes  
6      in the detectable property generate data evidencing each monomer incorporation producing a  
7      monomer sequence read out.

1      11.(previously presented)    The composition of claim 10, wherein the change in the fluorescence  
2      property results from a change in the conformation of the polymerizing agent from a first  
3      conformational state to a second conformational state and back again during each monomer  
4      incorporation.

1      12.(currently amended)      The composition of claim ~~10~~ 11, wherein the fluorescence property  
2      has a first detection propensity when the polymerizing agent is in the first conformational state and  
3      a second detection propensity when the polymerizing agent is in the ~~a~~ second conformational state.



20.(canceled)

21.(canceled)

22.(canceled)

22.(canceled)

23.(canceled)

24.(canceled)

1     25.(withdrawn)     A single molecule sequencing apparatus comprising a substrate having a first  
2     chamber in which at least one tagged polymerase is confined therein and a second chamber including  
3     tagged dNTPs and a channel interconnecting the chambers, where a detectable property of at least  
4     one tag undergoes a detectable change during a monomer incorporation cycle.

1     26.(withdrawn)     The apparatus of claims 24, further comprising a plurality of monomer  
2     chambers, one for each tagged dNTP.

1     27.(withdrawn)     A mutant Taq polymerase comprising native Taq polymerase with a cysteine  
2     residue replacement at a site selected from the group consisting of 513-518, 643, 647, 649 and 653-  
3     661 and mixtures or combinations thereof.

1     28.(withdrawn)     The polymerase of claim 27, wherein the cysteine residue includes a tag  
2     covalently bonded thereto through the SH group.

1     29.(withdrawn)     A system for retrieving stored information comprising:  
2             a unknown nucleotide sequence representing a data stream;  
3             a single-molecule sequencer including a polymerase having a tag associated therewith and  
4     monomers for the polymerase, each monomer having a tag associated therewith;  
5             an excitation source adapted to excite the at least one of the tags; and  
6             a detector adapted to detect a response from at least one of the tag,  
7             where the response changes during polymerization of a complementary sequence and the  
8     changes in response represent a content of the data stream.

1 30.(withdrawn) A system for determining sequence information from a single molecule  
2 comprising:

3 a unknown nucleotide sequence;  
4 a single-molecule sequencer comprising a polymerase having a tag associated therewith and  
5 monomers for the polymerase, each monomer having a tag associated therewith;  
6 a excitation source adapted to excite at least one of the tags; and  
7 a detector adapted to detect a response from at least one of the tags,  
8 where the response changes during polymerization of a complementary sequence and the  
9 changes in the response represent the identity of each nucleotide in the unknown sequence.

1 31.(withdrawn) A method for sequencing a molecular sequence comprising:

2 supplying an unknown sequence of nucleotides or nucleotide analogs to a single-molecule  
3 sequencer comprising a polymerase having a fluorescent donor covalently attached thereto and  
4 monomers for the polymerase, each monomer having a unique fluorescent acceptor covalently  
5 bonded thereto;

6 exciting the fluorescent donor with a light from an excitation light source;  
7 detecting emitted fluorescent light from the acceptor during a monomer incorporation cycle  
8 via a fluorescent light detector, where an intensity and/or frequency of the emitted light for the  
9 acceptors changes during each monomer incorporation cycle; and

10 converting the changes into an identity of each nucleotide or nucleotide analog in the  
11 unknown sequene.

1 32.(withdrawn) A method of sequencing an individual nucleic acid molecule or numerous  
2 individual molecules in parallel including the steps of:

3 immobilizing a member of the replication complex comprising a polymerase including a tag  
4 attached thereto, a primer or a template sufficiently spaced apart to allow resolution detection of each  
5 complex on a solid support;

6 incubating the replication complex with cooperatively-tagged nucleotides, each nucleotide  
7 including a unique tag at its gamma-phosphate, where each nucleotide can be individually detected;

8 detecting each nucleotide incorporated by the polymerase as the polymerase transitions  
9 between its open and closed form, which causes a change in a detectable property of at least one of

10 the tags or as the pyrophosphate group is released by the polymerase; and  
11 relating the changes in the detectable property to the sequence of nucleotides in an unknown  
12 nucleic acid sequence.

1 33.(withdrawn) A  $\gamma$ -phosphate modified nucleoside comprising  $\gamma$ -phosphate modified dATP,  
2 dCTP, dGTP and dTTP.

1 34.(withdrawn) A primer sequence or portion thereof selected from the group consisting of  
2 Sequence 1 through 29.

35.(canceled)

36.(canceled)

37.(canceled)

38.(canceled)

39.(canceled)

40.(canceled)

41.(canceled)

42.(canceled)

43.(canceled)

44.(canceled)

45.(canceled)

46.(canceled)

47.(canceled)

1 48.(canceled) A composition comprising a polymerizing agent including at least one molecular  
2 and/or atomic tag covalently bonded to a site on the polymerizing agent, where a fluorescence  
3 property of the tags undergoes a change before, during and/or after each of a sequence of monomer  
4 incorporations and where the changes in the fluorescence property generate data evidencing each  
5 monomer incorporation producing a monomer incorporation read out and where the polymerizing  
6 agent comprises a *Taq* DNA polymerase I having a tag covalently bonded to an amino acid site of

7 the *Taq* polymerase selected from the group consisting of 513-518, 643, 647, 649 and 653-661 and,  
8 where the tag comprises a fluorescent molecule.

1 49.(canceled) The composition of claim 48, wherein the fluorescence property has a first value  
2 when the polymerizing agent is in a first state and a second value when the polymerizing agent is in  
3 a second state, and where the polymerizing agent changes from the first state to the second state and  
4 back again during each monomer incorporation.